

Search #1 $R_2 = R_3 = H$

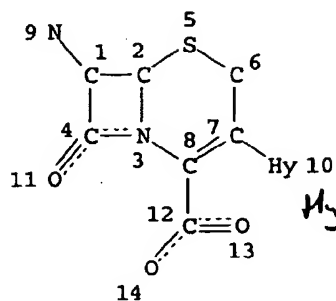
M. Berch; 10/006,279

Page 1

=> d que 17

L1

STR



Initial structure allows R_2 & R_3 to be anything.



NODE ATTRIBUTES:

DEFAULT MLEVEL IS ATOM
GGCAT IS MCY SAT AT 10
DEFAULT ECLEVEL IS LIMITED
ECOUNT IS E4 C E1 O AT 10

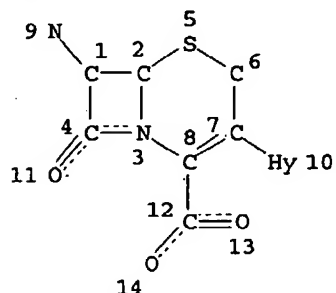
GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 14

STEREO ATTRIBUTES: NONE

L3 148 SEA FILE=REGISTRY SSS FUL L1
L4 STR

Initial search produces subset of 148 hits



this structure limits $R_2 = R_3 = H$

NODE ATTRIBUTES:

CONNECT IS E1 RC AT 9
CONNECT IS E1 RC AT 14
DEFAULT MLEVEL IS ATOM
GGCAT IS MCY SAT AT 10
DEFAULT ECLEVEL IS LIMITED
ECOUNT IS E4 C E1 O AT 10

Connectivity set to Exactly 1 @ 9&14 - remaining open positions can only be H (otherwise connect is >1).

GRAPH ATTRIBUTES:

RSPEC I
NUMBER OF NODES IS 14

Generic hy @ 10 must have Exactly 4 Carbons (E4C) and exactly 1 oxygen (E1O). It must be monocyclic (MCY) and saturated (SAT).

STEREO ATTRIBUTES: NONE

L6 1 SEA FILE=REGISTRY SUB=L3 SSS FUL L4
L7 1 SEA FILE=CAPLUS ABB=ON PLU=ON L6

subset search of subset L3 using str L4 produces 1 hit in Registry. Searching CAPLUS with Registry answer set also produces 1 hit.

=> D IBIB ABS HITSTR

Searched by Thom Larson, STIC, 308-7309

Point of Contact:
Thomas G. Larson, Ph.D.
703-308-7309
CM1, Rm. 6 B 01

L7 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:449688 CAPLUS

DOCUMENT NUMBER: 137:33161

TITLE: Coupling process and intermediates useful for preparing cephalosporins

INVENTOR(S): Colberg, Juan Carlos; Donadelli, Alessandro; Fogliato, Giovanni; Zenoni, Maurizio

PATENT ASSIGNEE(S): Pfizer Products Inc., USA

SOURCE: PCT Int. Appl., 33 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

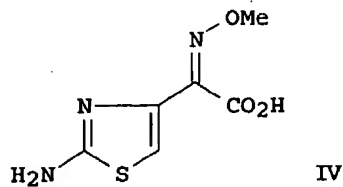
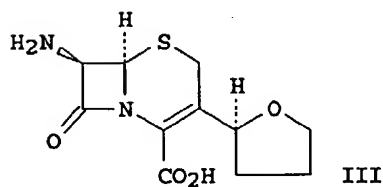
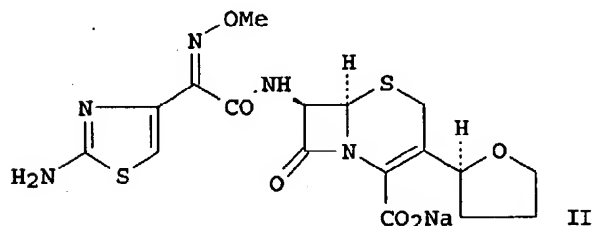
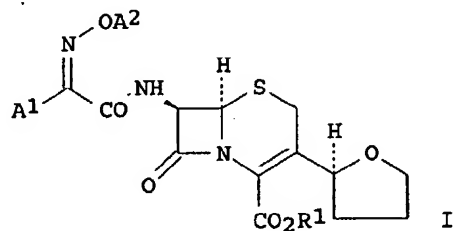
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002046198	A1	20020613	WO 2001-IB2225	20011122
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.:

US 2000-251014P P 20001204

OTHER SOURCE(S): MARPAT 137:33161

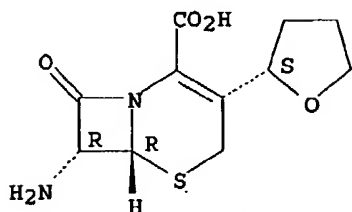
GI



*Looks like
your applic-
ants' work.*

- AB This invention relates to a novel process for the prepn. of 3-cyclic-ether-substituted cephalosporins, such as I [CO₂R₁ = carboxylic acid or a carboxylate salt; A₁ = aryl, heteroaryl, heterocyclyl; A₂ = H, alkyl, cycloalkyl, aryl, etc.], via amidation reactions. Thus, cephalosporin II was prepd. in 80% yield by amidation of amine III with the acid anhydride of acid IV using O,O-di-Et hydrogenphosphorothioate in a Me₂CO/H₂O soln.
- IT 436100-71-9P
RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(process for the prepn. of intermediates via amidation which are useful for prepg. cephalosporins)
- RN 436100-71-9 CAPLUS
- CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-amino-8-oxo-3-[(2S)-tetrahydro-2-furanyl]-, (6R,7R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT:

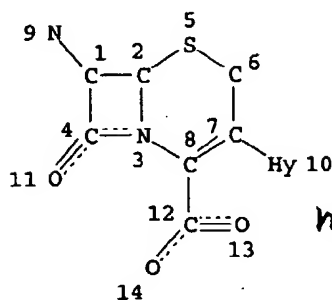
6

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

$\Rightarrow d \text{ que } 16$

L1

STR



Initial search same as
in part 1 - R_2 & R_3
left open

NODE ATTRIBUTES:

DEFAULT MLEVEL IS ATOM

GGCAT IS MCY SAT AT 10

DEFAULT ECLEVEL IS LIMITED

ECOUNT IS E4 C E1 O AT 10

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

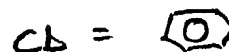
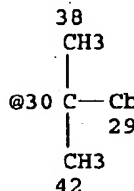
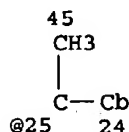
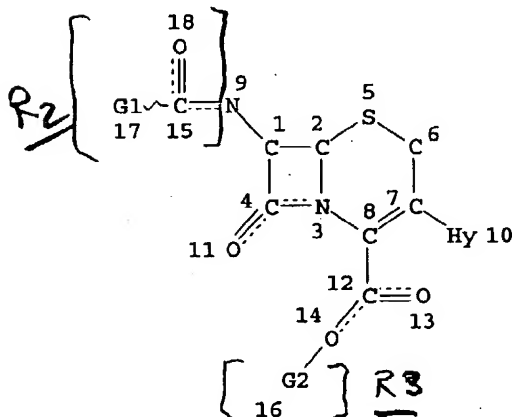
NUMBER OF NODES IS 14

STEREO ATTRIBUTES: NONE

L2 148 SEA FILE=REGISTRY SSS FUL L1

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100	100

Final search results.



VAR G1=H/C/N/O

VAR G2=19/25/30 - R3

NODE ATTRIBUTES:

CONNECT IS E2 RC AT 19

CONNECT IS E1 RC AT 25

DEFAULT MLEVEL IS ATOM

GGCAT IS MCY SAT AT 10

GGCAT IS MCY UNS AT 20

GGCAT	IS	MCY	UNS	AT	24
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GGCAT	IS	MCY	UNS	AT	29
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DEFAULT ECLEVEL IS LIMITED

ECOUNT IS E4 C E1 0 AT 10

E6 C AT 20

ECOUNT	IS	E6	C	AT	24
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ECOUNT 15 00 C A1 21

includes substructures with attachment points \oplus [#]19, 25, 230.

- open bonds on 19 are limited to H only.
- open bond on 25 is limited to H only.
- generic by group @ 10 is limited to being monocyclic and unsaturated.

generic group $CB @ 20, 24, \geq 29$ is limited to being monocyclic and unsaturated.

¹⁰ — $H_g @ 10$ has exactly 4C and 10.

- ch @ 20, 24, 30 has exactly 6 C

ECOUNT IS E6 C AT 29

GRAPH ATTRIBUTES:

RSPEC I

NUMBER OF NODES IS 27

STEREO ATTRIBUTES: NONE

L5 32 SEA FILE=REGISTRY SUB=L2 SSS FUL L3
 L6 7 SEA FILE=HCAPLUS ABB=ON PLU=ON L5

*search subset L2 with
 structure L3 produces
 32 hits. Searching
 Hcaplus with Registry
 answer set produces
 7 hits,*

=> D IBIB ABS HITSTR 1-7

L6 ANSWER 1 OF 7 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:104661 HCAPLUS

DOCUMENT NUMBER: 136:151036

TITLE: Process for the preparation of cephalosporin compounds and their intermediates

INVENTOR(S): Burton, George; Best, Desmond John; Gasson, Brian Charles; Osborne, Neal Frederick; Walker, Graham

PATENT ASSIGNEE(S): Pfizer Inc., USA

SOURCE: Eur. Pat. Appl., 22 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1178049	A1	20020206	EP 2001-306325	20010723
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
US 2002058806	A1	20020516	US 2001-918152	20010730
JP 2002105083	A2	20020410	JP 2001-233551	20010801
PRIORITY APPLN. INFO.:			GB 2000-19124	A 20000803

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB A process for prep. cephalosporins I (R1 = H, OMe, formamido; R2 = acyl; CO2R3 = carboxy group or CO2- or readily removable carboxy protecting group; R4 = H, or up to four substituents from alkyl, alkenyl, alkynyl, alkoxy, halogen, amino, alkyl(acyl)amino, CO2R, CONR2, SO2NR2 (R = H, C1-6 alkyl), aryl, heterocycle, etc.; X = S, SO, SO2, O, CH2; m = 1-2; dotted lines indicate a 2- or 3-cephem system) was accomplished via the cyclization of II. Thus the 3-(R and S)-tetrahydrofuran-2-yl-2-em compds. III were prep. and the S isomer was converted to the 3-(S)-tetrahydrofuran-2-yl-3-em III in several steps.

IT 141194-60-7P

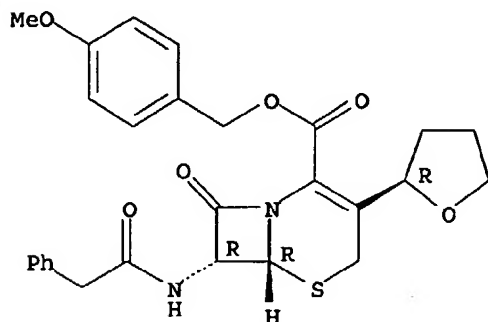
RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (process for prep. cephalosporin compds. and their intermediates)

RN 141194-60-7 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
 8-oxo-7-[(phenylacetyl)amino]-3-[(2R)-tetrahydro-2-furanyl]-,

(4-methoxyphenyl)methyl ester, (6R,7R) - (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 2 OF 7 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1997:140924 HCAPLUS

DOCUMENT NUMBER: 126:144046

TITLE: Beta-lactam preparation

INVENTOR(S): Harris, Michael Anthony; Saunders, Richard Neville

PATENT ASSIGNEE(S): Pfizer Limited, UK

SOURCE: Brit. UK Pat. Appl., 15 pp.

CODEN: BAXXDU

DOCUMENT TYPE: Patent

LANGUAGE: English

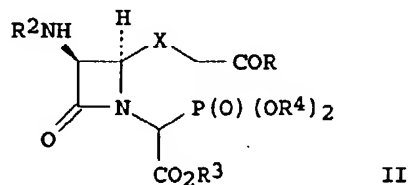
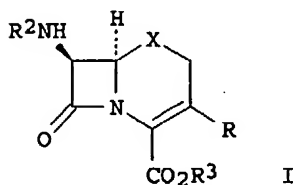
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
GB 2300856	A1	19961120	GB 1995-10126	19950516

OTHER SOURCE(S): CASREACT 126:144046; MARPAT 126:144046

GI



AB Title compds. I [R = substituent; R1 = H, OMe, NHCHO; R2 = acyl; CO2R3 = CO2H, CO2-; R3 = protecting group; X = S, SO, SO2, O, CH2] are prepd. by base-induced cyclization of an azetidinone II [R4 = alkyl, aryl]. II are prepd. from the halide and P(OR4)3. Thus, 4-methoxybenzyl (2RS)-2-hydroxy-2-[(3R) (4R)-3-phenylacetamido-4-[(RS)-2-tetrahydrofuryl]carbonylmethylthio]azetidin-2-on-1-ylacetate was converted to the chloride and then to the phosphonate which was cyclized with NaH in PhMe to give 50% I [R = (RS)-2-tetrahydrofuryl, R1 = H, R2 = PhCH2CO, R3 = 4-MeC6H4CH2].

IT 141061-21-4P

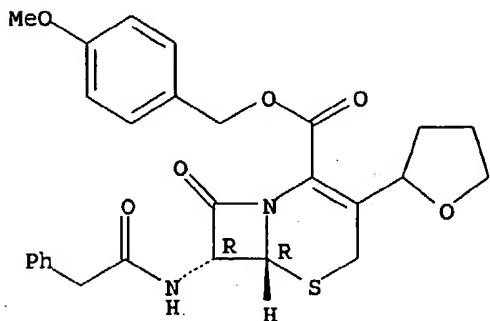
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(prepn. of cepems by cyclization of azetidylphosphonoacetates with base)

RN 141061-21-4 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid, 8-oxo-7-[(phenylacetyl)amino]-3-(tetrahydro-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-(6.alpha.,7.beta.)]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L6 ANSWER 3 OF 7 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1997:1373 HCAPLUS

DOCUMENT NUMBER: 126:89179

TITLE: Transformation of penicillins into 3-substituted .DELTA.3-cephems through addition/cyclization of allenecarboxylates

AUTHOR(S): Tanaka, Hideo; Sumida, Shin-ichi; Kameyama, Yutaka; Sorajo, Koichi; Wada, Isao; Torii, Sigeru

CORPORATE SOURCE: Faculty of Engineering, Okayama University, Okayama, 700, Japan

SOURCE: Bulletin of the Chemical Society of Japan (1996), 69(12), 3651-3658

CODEN: BCSJA8; ISSN: 0009-2673

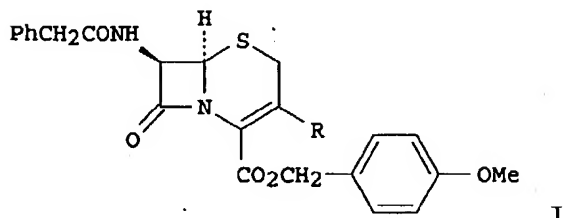
PUBLISHER: Nippon Kagakkai

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 126:89179

GI



I

AB A straightforward synthesis of .DELTA.3-cephems I [R = morpholino,

pyrrolidino; N3, 5-methyl-1,3,4-thiadiazolyl-2-thio, SO₂Ph] was performed successfully by a sequential addn./cyclization reaction of the allenecarboxylate derived from penicillin. The addn./cyclization reaction proceeded smoothly upon treatment of the allenecarboxylate with the nucleophiles. Reaction of the allenecarboxylate with lithium chloride in NMP (N-methyl-2-pyrrolidone) in the presence of aluminum chloride afforded I [R = Cl], while without AlCl₃ I [R = SO₂Ph] was the main product. I [R = arylthio] were similarly prepd.

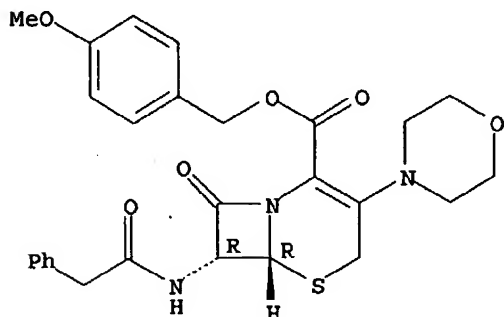
IT 139472-61-0P

RL: SPN (Synthetic preparation); PREP (Preparation)
(transformation of penicillins into 3-substituted .DELTA.3-cephems
through addn./cyclization of allenecarboxylates)

RN 139472-61-0 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
3-(4-morpholinyl)-8-oxo-7-[(phenylacetyl)amino]-, (4-methoxyphenyl)methyl
ester, (6R-trans)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L6 ANSWER 4 OF 7 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1996:456093 HCAPLUS

DOCUMENT NUMBER: 125:114393

TITLE: Process for the preparation of cephalosporins and
analogs

INVENTOR(S): Burton, George; Naylor, Antoinette

PATENT ASSIGNEE(S): Pfizer Inc., USA

SOURCE: PCT Int. Appl., 29 pp.

CODEN: PIXXD2

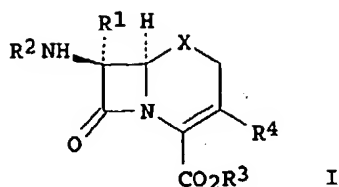
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9617847	A1	19960613	WO 1995-GB2783	19951129
W: JP, US				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
PRIORITY APPLN. INFO.:			GB 1994-24847	19941209
OTHER SOURCE(S):			CASREACT 125:114393; MARPAT 125:114393	
GI				



AB Cephalosporins I [X = S, SO, SO₂, O, CH₂; R₁ = H, OMe, NHCHO; R₂ = acyl; R₃ = in vivo hydrolyzable ester group; R₄ = (un)substituted tetrahydrofuryl, tetrahydropyranyl] are prepd. by reaction of the corresponding carboxylic acid with R₃Y [Y = halide] in the presence of an aq. phase contg. a base and a phase transfer catalyst. Subsequent removal of protecting groups, conversion of groups X and R₂ and salt formation may be carried out. Thus, 4-methoxybenzyl (6R,7R)-7-phenylacetamido-3-[(S)-2-tetrahydrofuryl]cephem-4-carboxylate was treated with Me₃CCO₂CH₂I, followed by deacylation and reacylation to give pivaloyloxymethyl (6R,7R)-7-[2-(2-amino-4-thiazolyl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-2-tetrahydrofuryl]cephem-4-carboxylate.

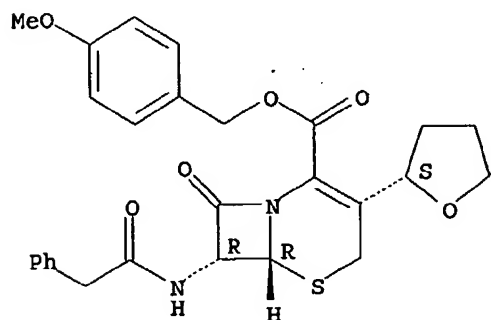
IT 141194-63-0

RL: RCT (Reactant); RACT (Reactant or reagent)
(prepn. and transacylation of cephalosporin esters)

RN 141194-63-0 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-(tetrahydro-2-furanyl)-,
(4-methoxyphenyl)methyl ester, [6R-[3(S*),6.alpha.,7.beta.]]- (9CI) (CA
INDEX NAME)

Absolute stereochemistry.



L6 ANSWER 5 OF 7 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1993:38686 HCAPLUS

DOCUMENT NUMBER: 118:38686

TITLE: Process for preparing cephem derivatives

INVENTOR(S): Torii, Sigeru; Tanaka, Hideo; Taniguchi, Masatoshi;

Sasaoka, Michio; Shiroi, Takashi; Kameyama, Yutaka

PATENT ASSIGNEE(S): Otsuka Kagaku K. K., Japan

SOURCE: Eur. Pat. Appl., 13 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

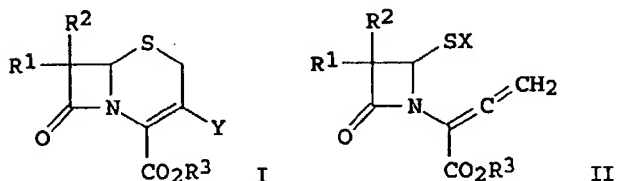
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

Searched by Thom Larson, STIC, 308-7309

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 507124	A2	19921007	EP 1992-104202	19920311
EP 507124	A3	19921209		
EP 507124	B1	19961227		
R: DE, FR, GB, IT				
JP 04282387	A2	19921007	JP 1991-72450	19910311
JP 3195371	B2	20010806		
US 5204458	A	19930420	US 1992-849160	19920310
PRIORITY APPLN. INFO.:			JP 1991-72450	A 19910311
OTHER SOURCE(S):	CASREACT 118:38686; MARPAT 118:38686			
GI				



AB Title compds. [I; R1 = (protected) amino; R2 = H, alkoxy; R3 = H, carboxy-protective group; Y = residue of a nucleophile] were prepd. by cyclization of azetidinylenethenylideneacetates II [R1-R3 as above; X = SO2R4, SR4; R4 = aryl, (substituted) N-contg. heteroaryl] with a nucleophile. Thus, II [R1 = PhCH2CONH, R2 = H, R3 = CH2C6H4(OMe)-4, X = SO2Ph] was stirred 1 h with morpholine in DMF contg. CaCl2 to give 87% I (R1-R3 unchanged, Y = morpholino).

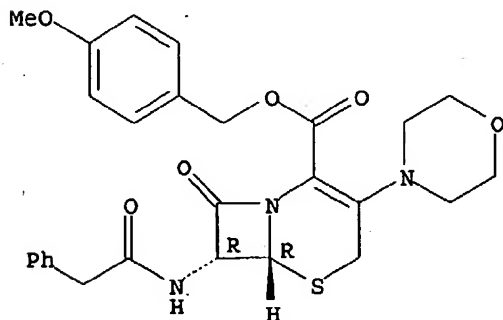
IT 139472-61-0P

RL: SPN (Synthetic preparation); PREP (Preparation)
(prepn. of, method for)

RN 139472-61-0 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
3-(4-morpholinyl)-8-oxo-7-[(phenylacetyl)amino]-, (4-methoxyphenyl)methyl
ester, (6R-trans)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L6 ANSWER 6 OF 7 HCAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 1992:255397 HCAPLUS

Searched by Thom Larson, STIC, 308-7309

DOCUMENT NUMBER: 116:255397
 TITLE: Preparation of 3-tetrahydrofurylcephem-3-carboxylates and analogs as antibiotics
 INVENTOR(S): Bateson, John Hargreaves; Burton, George; Fell, Stephen Christopher Martin
 PATENT ASSIGNEE(S): Beecham Group PLC, UK
 SOURCE: PCT Int. Appl., 147 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9201696	A1	19920206	WO 1991-GB1228	19910722
W: AU, CA, CS, FI, HU, JP, KR, NO, PL, US				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LU, NL, SE				
CA 2087967	AA	19920125	CA 1991-2087967	19910722
AU 9182224	A1	19920218	AU 1991-82224	19910722
AU 648329	B2	19940421		
ZA 9105725	A	19920624	ZA 1991-5725	19910722
EP 540609	A1	19930512	EP 1991-913583	19910722
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE				
HU 63628	A2	19930928	HU 1993-177	19910722
JP 05509305	T2	19931222	JP 1991-512368	19910722
JP 2851428	B2	19990127		
AT 185567	E	19991015	AT 1991-913583	19910722
ES 2137162	T3	19991216	ES 1991-913583	19910722
CN 1060469	A	19920422	CN 1991-105783	19910724
CN 1061046	B	20010124		
NO 9300226	A	19930323	NO 1993-226	19930122
US 6020329	A	20000201	US 1997-958864	19971020
CN 1223859	A	19990728	CN 1998-122407	19981114
US 6001997	A	19991214	US 1999-228138	19990111
US 6077952	A	20000620	US 1999-327667	19990608
PRIORITY APPLN. INFO.:				
			GB 1990-16189	A 19900724
			GB 1991-9540	A 19910502
			WO 1991-GB1228	A 19910722
			US 1993-934667	B1 19930122
			US 1995-470786	B1 19950606
			US 1997-958864	A1 19971020
			US 1999-228138	A1 19990111

OTHER SOURCE(S): MARPAT 116:255397

GI For diagram(s), see printed CA Issue.

AB Title compds. (I; R1 = H, MeO, HCONH; R2 = acyl; R3 = H, neg. charge, carboxy-protective group; R4 = .ltoreq.4 substituents selected from alkyl, alkenyl, OH, halo, alkoxy, etc.; X = O, CH2, SOn; n= 0-2; m = 1, 2) were prep'd. Thus, Na 2-(2-tritylaminothiazol-4-yl)-2-(Z)-trityloxyiminoacetate was condensed with tert-butyl (6R, 7R)-7-amino-3-[(R)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate to give, after deprotection, (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-hydroxyiminoacetamido]-3-[(RS)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylic acid which had MIC of 0.50 and 0.25 .mu.g/mL against Escherichia coli (NCTC 1048) and Staphylococcus aureus (Oxford), resp.

IT 141061-21-4P 141061-24-7P 141061-25-8P
 141072-52-8P 141072-54-0P 141072-55-1P
 141072-56-2P 141072-58-4P 141072-59-5P
 141072-60-8P 141072-67-5P 141072-71-1P
 141072-73-3P 141072-74-4P 141072-78-8P

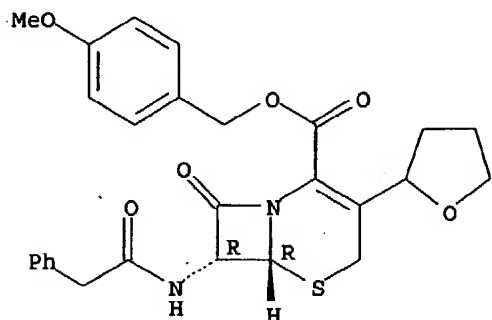
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 141072-87-9P 141072-94-8P 141072-95-9P
 141073-02-1P 141073-04-3P 141073-15-6P
 141194-60-7P 141194-63-0P 141194-67-4P
 141194-73-2P 141194-77-6P 141194-83-4P
 141194-85-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)
 (prepn. and reaction of, in prepn. of antibiotics)

RN 141061-21-4 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
 8-oxo-7-[(phenylacetyl)amino]-3-(tetrahydro-2-furanyl)-,
 (4-methoxyphenyl)methyl ester, [6R-(6.alpha.,7.beta.)]- (9CI) (CA INDEX
 NAME)

Absolute stereochemistry.

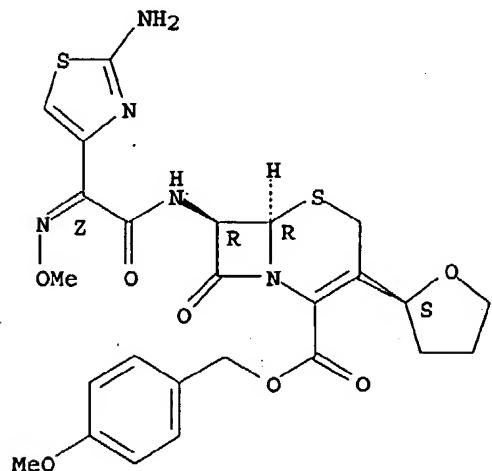


RN 141061-24-7 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
 7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-2-
 furanyl)-, (4-methoxyphenyl)methyl ester, [6R-[3(S*),6.alpha.,7.beta.(Z)]]-
 (9CI) (CA INDEX NAME)

Absolute stereochemistry.

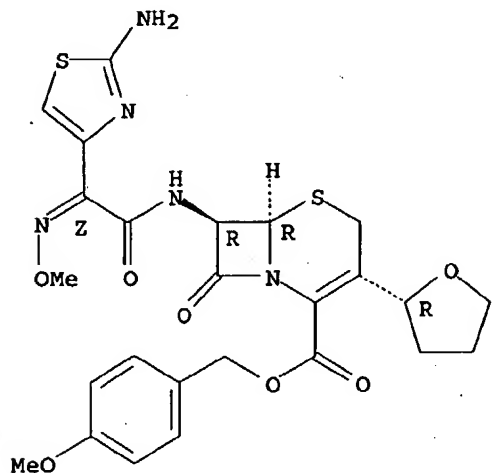
Double bond geometry as shown.



RN 141061-25-8 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-[3(R*),6.alpha.,7.beta.(Z)]]-(9CI) (CA INDEX NAME)

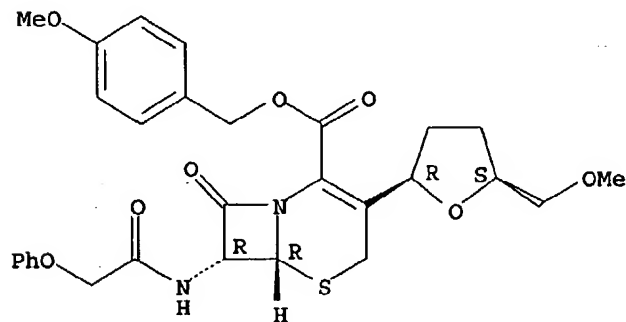
Absolute stereochemistry.
Double bond geometry as shown.



RN 141072-52-8 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenoxyacetyl)amino]-3-[tetrahydro-5-(methoxymethyl)-2-furanyl]-, (4-methoxyphenyl)methyl ester, [6R-[3(2R*,5S*),6.alpha.,7.beta.]]-(9CI) (CA INDEX NAME)

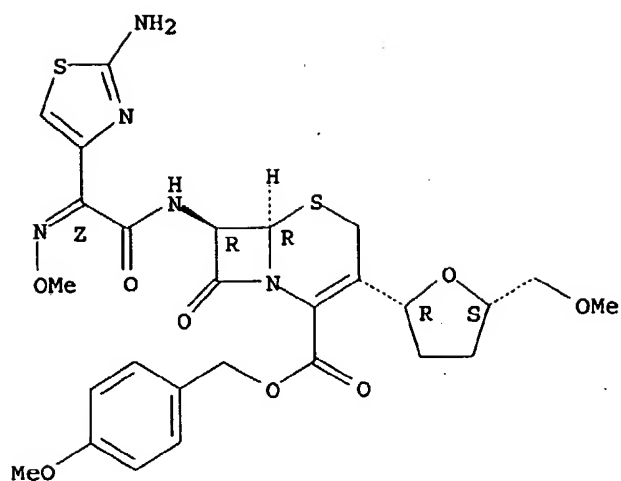
Absolute stereochemistry.



RN 141072-54-0 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-[tetrahydro-5-(methoxymethyl)-2-furanyl]-, (4-methoxyphenyl)methyl ester, [6R-[3(2R*,5S*),6.alpha.,7.beta.(Z)]]-(9CI) (CA INDEX NAME)

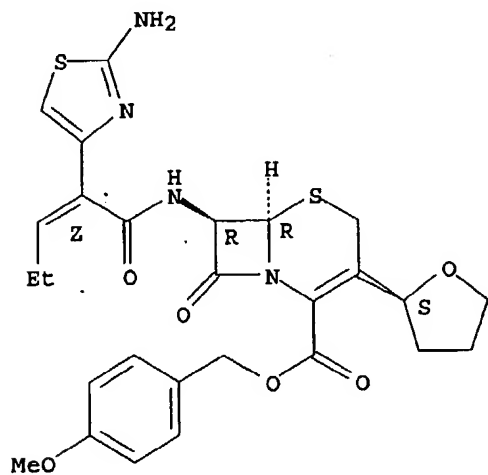
Absolute stereochemistry.
Double bond geometry as shown.



RN 141072-55-1 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[2-(2-amino-4-thiazolyl)-1-oxo-2-pentenyl]amino]-8-oxo-3-(tetrahydro-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-[3(S*),6.alpha.,7.beta.(Z)]]-(9CI) (CA INDEX NAME)

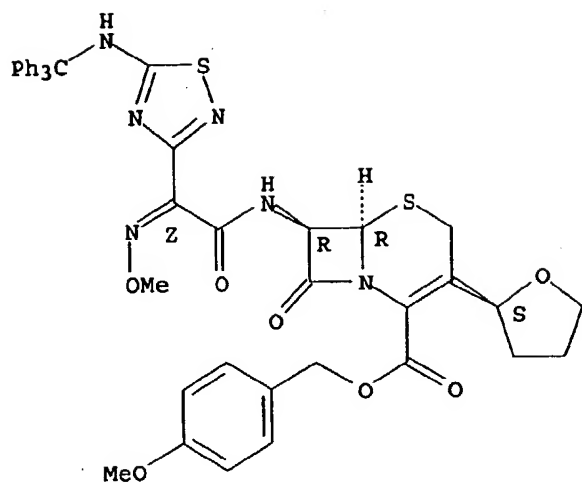
Absolute stereochemistry.
Double bond geometry as shown.



RN 141072-56-2 HCAPLUS

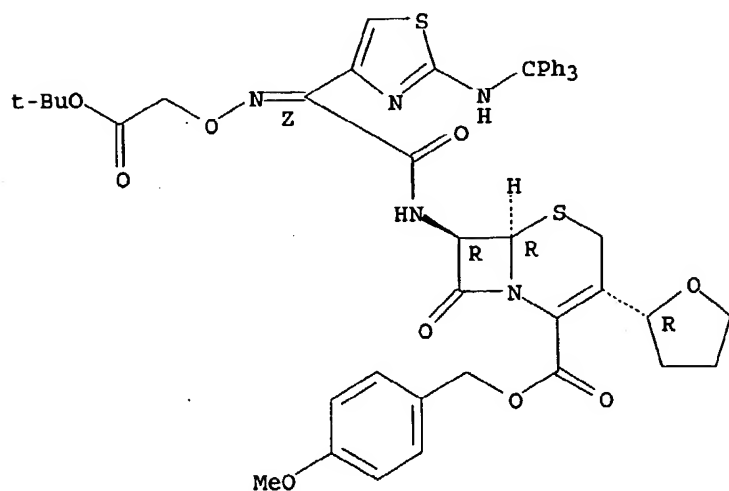
CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(methoxyimino) [5-[(triphenylmethyl)amino]-1,2,4-thiadiazol-3-yl]acetyl]amino]-8-oxo-3-(tetrahydro-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-[3(S*),6.alpha.,7.beta.(Z)]]-(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.



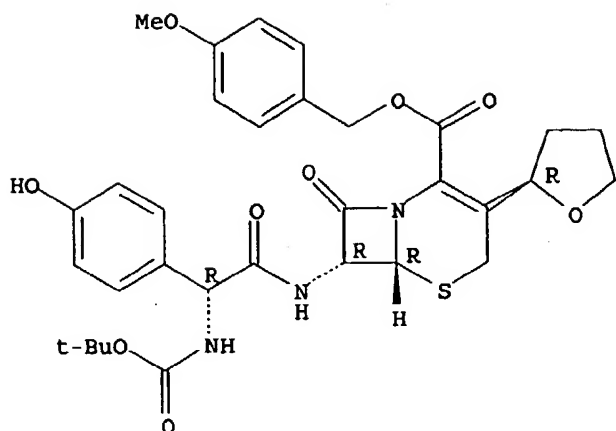
RN 141072-58-4 HCAPLUS
 CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
 7-[[[2-(1,1-dimethylethoxy)-2-oxoethoxy]imino][2-[(triphenylmethyl)amino]-
 4-thiazolyl]acetyl]amino]-8-oxo-3-(tetrahydro-2-furanyl)-,
 (4-methoxyphenyl)methyl ester, [6R-[3(R*),6.alpha.,7.beta.(Z)]]- (9CI)
 (CA INDEX NAME)

Absolute stereochemistry.
 Double bond geometry as shown.



RN 141072-59-5 HCAPLUS
 CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
 7-[[[[(1,1-dimethylethoxy)carbonyl]amino](4-hydroxyphenyl)acetyl]amino]-8-
 oxo-3-(tetrahydro-2-furanyl)-, (4-methoxyphenyl)methyl ester,
 [6R-[3(R*),6.alpha.,7.beta.(R*)]]- (9CI) (CA INDEX NAME)

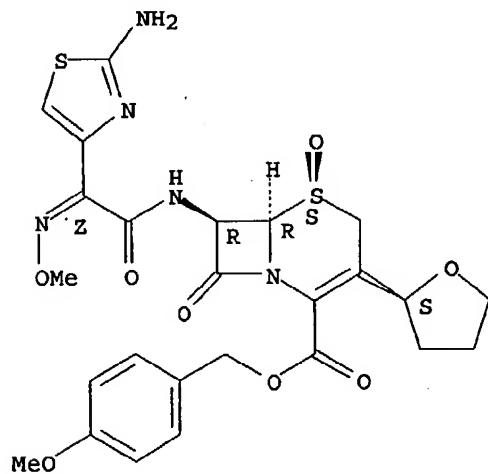
Absolute stereochemistry.



RN 141072-60-8 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-2-furanyl)-, (4-methoxyphenyl)methyl ester, 5-oxide, [5S-
[3(R*),5.alpha.,6.beta.,7.alpha.(Z)]]- (9CI) (CA INDEX NAME)

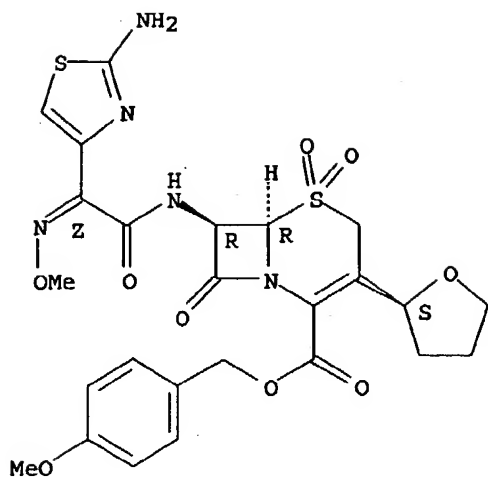
Absolute stereochemistry.
Double bond geometry as shown.



RN 141072-67-5 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-2-furanyl)-, (4-methoxyphenyl)methyl ester, 5,5-dioxide,
[6R-[3(S*),6.alpha.,7.beta.(Z)]]- (9CI) (CA INDEX NAME)

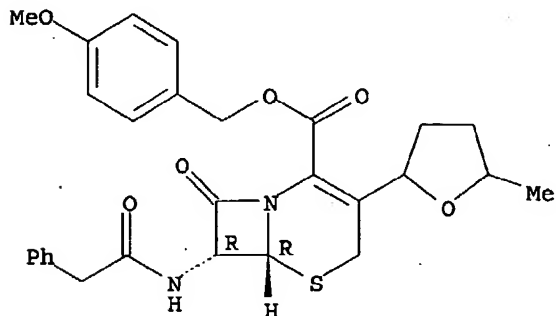
Absolute stereochemistry.
Double bond geometry as shown.



RN 141072-71-1 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-(tetrahydro-5-methyl-2-furanyl)-,
(4-methoxyphenyl)methyl ester, [6R-(6.alpha.,7.beta.)]- (9CI) (CA INDEX
NAME)

Absolute stereochemistry.

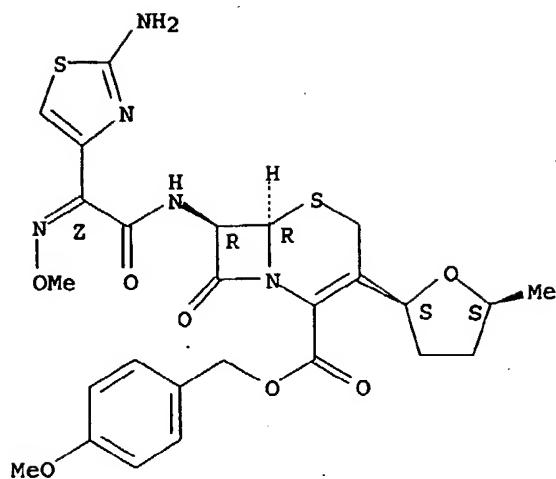


RN 141072-73-3 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-5-
methyl-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-
[3(2S*,5S*),6.alpha.,7.beta.(Z)]]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

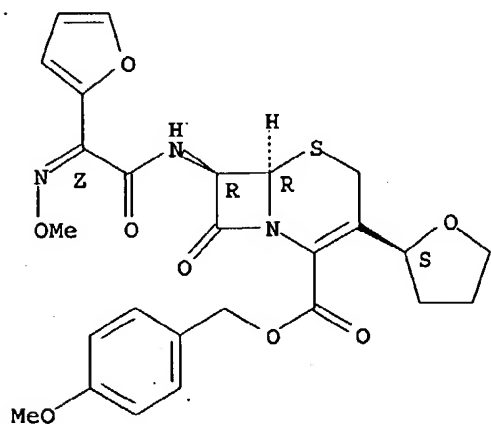
Double bond geometry as shown.



RN 141072-74-4 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[2-furanyl(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-2-furanyl)-,
(4-methoxyphenyl)methyl ester, [6R-[3(S*),6.alpha.,7.beta.(Z)]]- (9CI)
(CA INDEX NAME)

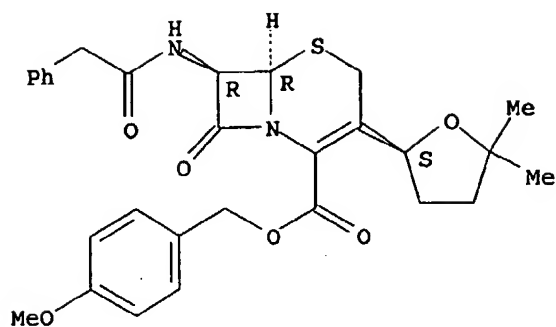
Absolute stereochemistry.
Double bond geometry as shown.



RN 141072-78-8 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-(tetrahydro-5,5-dimethyl-2-furanyl)-,
(4-methoxyphenyl)methyl ester, [6R-[3(S*),6.alpha.,7.beta.]]- (9CI) (CA
INDEX NAME)

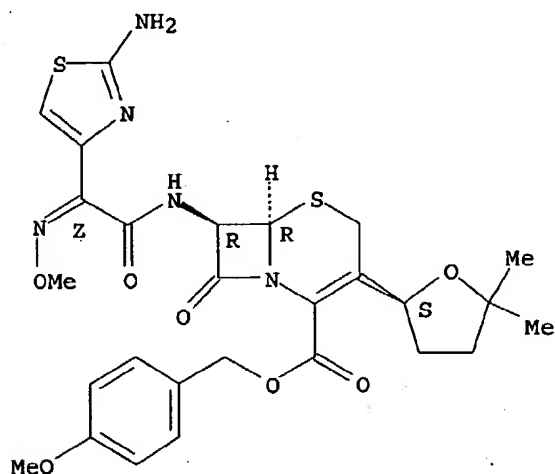
Absolute stereochemistry.



RN 141072-80-2 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-
5,5-dimethyl-2-furanyl)-, (4-methoxyphenyl)methyl ester,
[6R-[3(S*),6.alpha.,7.beta.(Z)]]- (9CI) (CA INDEX NAME)

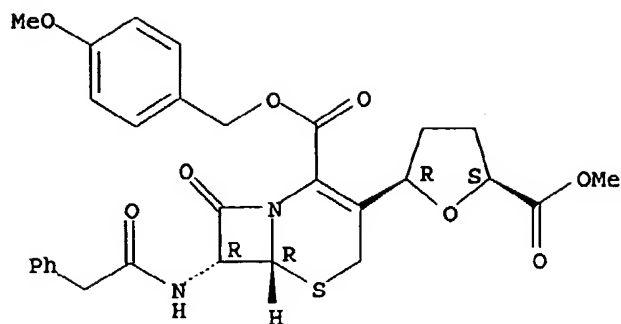
Absolute stereochemistry.
Double bond geometry as shown.



RN 141072-85-7 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-[tetrahydro-5-(methoxycarbonyl)-2-furanyl]-
, (4-methoxyphenyl)methyl ester, [6R-[3(2R*,5S*),6.alpha.,7.beta.]]- (9CI)
(CA INDEX NAME)

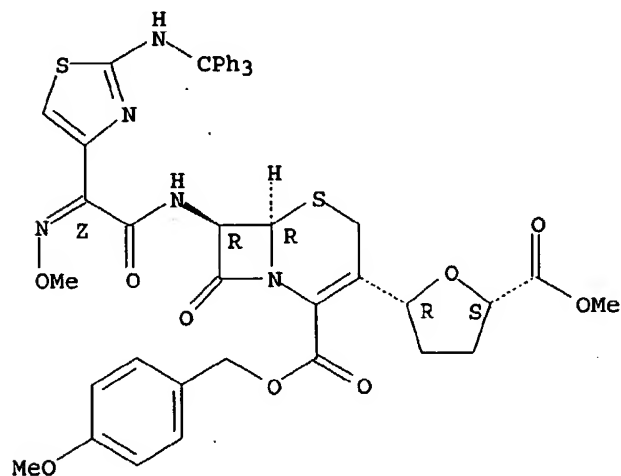
Absolute stereochemistry.



RN 141072-86-8 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(methoxyimino) [2-[(triphenylmethyl)amino]-4-thiazolyl]acetyl]amino]-8-
oxo-3-[tetrahydro-5-(methoxycarbonyl)-2-furanyl]-, (4-methoxyphenyl)methyl
ester, [6R-[3(2R*,5S*),6.alpha.,7.beta.(Z)]]- (9CI) (CA INDEX NAME)

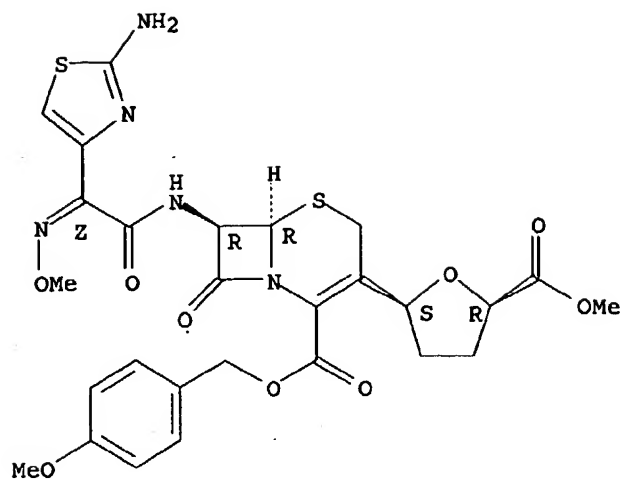
Absolute stereochemistry.
Double bond geometry as shown.



RN 141072-87-9 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl) (methoxyimino)acetyl]amino]-8-oxo-3-[tetrahydro-5-
(methoxycarbonyl)-2-furanyl]-, (4-methoxyphenyl)methyl ester,
[6R-[3(2S*,5R*),6.alpha.,7.beta.(Z)]]- (9CI) (CA INDEX NAME)

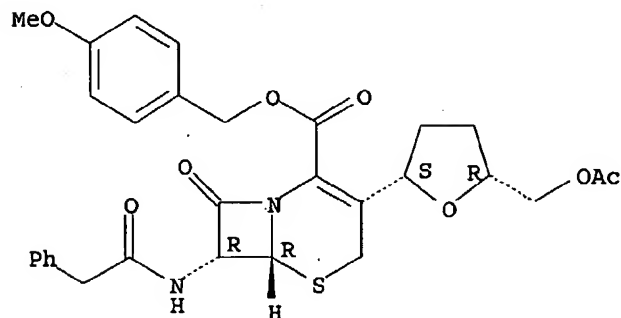
Absolute stereochemistry.
Double bond geometry as shown.



RN 141072-94-8 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
3-[5-[(acetyloxy)methyl]tetrahydro-2-furanyl]-8-oxo-7-
[(phenylacetyl)amino]-, (4-methoxyphenyl)methyl ester,
[6R-[3(2S*,5R*),6.alpha.,7.beta.]]- (9CI) (CA INDEX NAME)

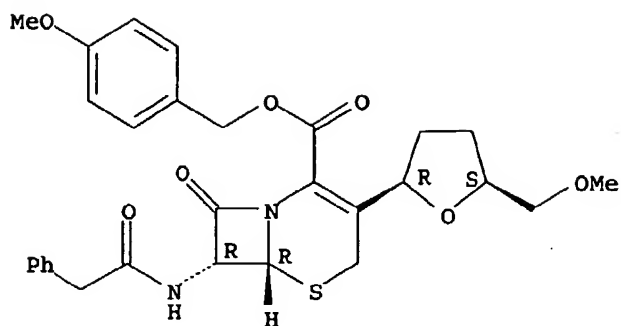
Absolute stereochemistry.



RN 141072-95-9 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-[tetrahydro-5-(methoxymethyl)-2-furanyl]-,
(4-methoxyphenyl)methyl ester, [6R-[3(2R*,5S*),6.alpha.,7.beta.]]- (9CI)
(CA INDEX NAME)

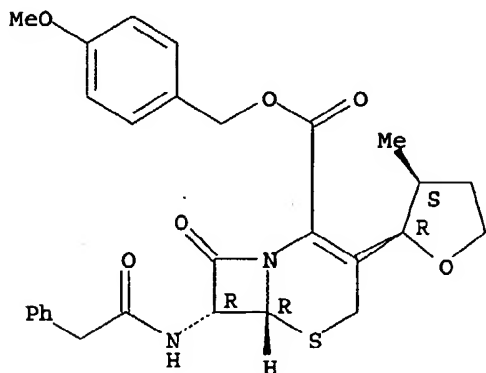
Absolute stereochemistry.



RN 141073-02-1 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-(tetrahydro-3-methyl-2-furanyl)-,
(4-methoxyphenyl)methyl ester, [6R-[3(2R*,3S*),6.alpha.,7.beta.]]- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.

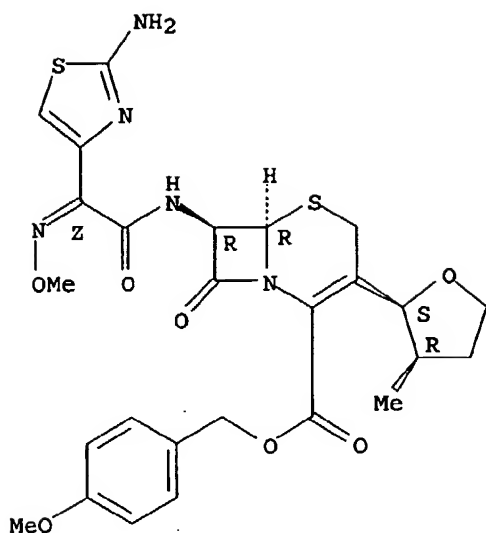


RN 141073-04-3 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-3-
methyl-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-
[3(2S*,3R*),6.alpha.,7.beta.(Z)]]- (9CI) (CA INDEX NAME)

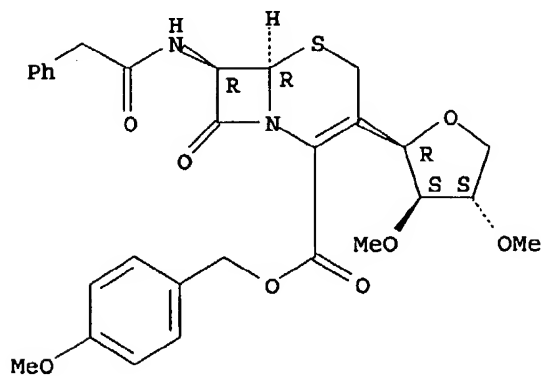
Absolute stereochemistry.

Double bond geometry as shown.



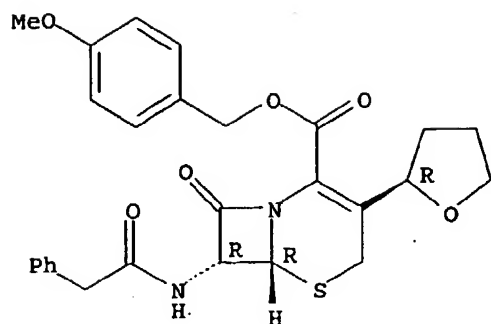
RN 141073-15-6 HCAPLUS
 CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
 8-oxo-7-[(phenylacetyl)amino]-3-(tetrahydro-3,4-dimethoxy-2-furanyl)-,
 (4-methoxyphenyl)methyl ester, [2R-[2.alpha.(6R*,7R*),3.alpha.,4.beta.]]-
 (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 141194-60-7 HCAPLUS
 CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
 8-oxo-7-[(phenylacetyl)amino]-3-[(2R)-tetrahydro-2-furanyl]-,
 (4-methoxyphenyl)methyl ester, (6R,7R)- (9CI) (CA INDEX NAME)

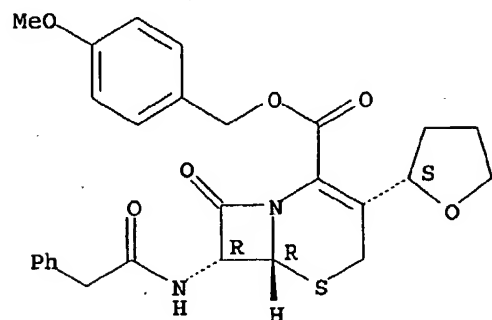
Absolute stereochemistry.



RN 141194-63-0 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-(tetrahydro-2-furanyl)-,
(4-methoxyphenyl)methyl ester, [6R-[3(S*),6.alpha.,7.beta.]]- (9CI) (CA
INDEX NAME)

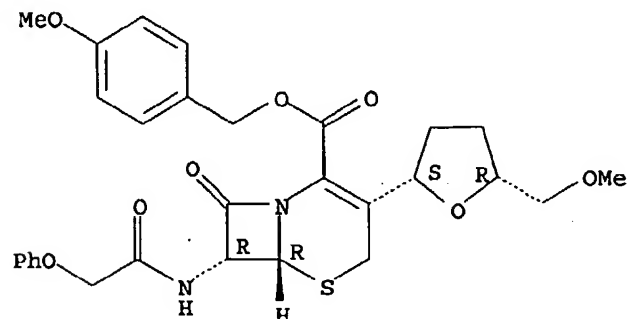
Absolute stereochemistry.



RN 141194-67-4 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenoxyacetyl)amino]-3-[tetrahydro-5-(methoxymethyl)-2-furanyl]-,
(4-methoxyphenyl)methyl ester, [6R-[3(2S*,5R*),6.alpha.,7.beta.]]- (9CI)
(CA INDEX NAME)

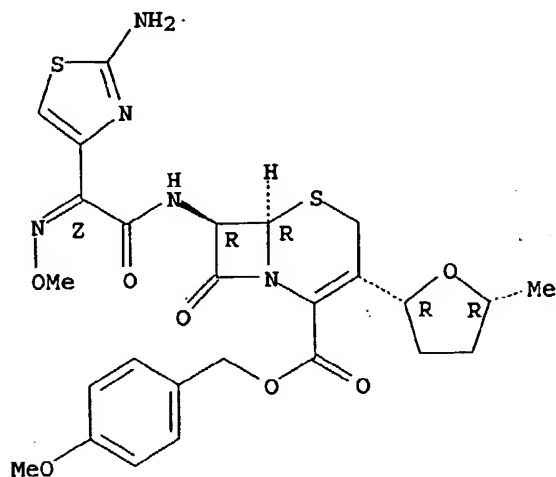
Absolute stereochemistry.



RN 141194-73-2 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-5-methyl-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-[3(2R*,5R*),6.alpha.,7.beta.(Z)]]- (9CI) (CA INDEX NAME)

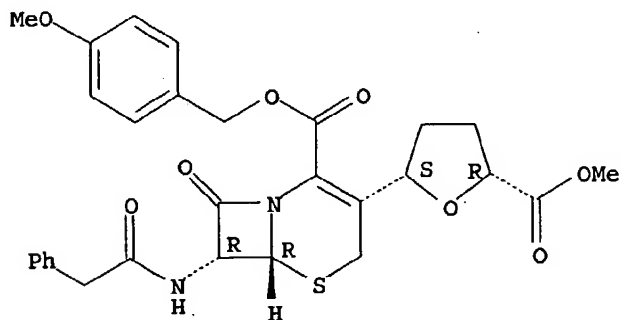
Absolute stereochemistry.
Double bond geometry as shown.



RN 141194-77-6 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-[(tetrahydro-5-(methoxycarbonyl)-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-[3(2S*,5R*),6.alpha.,7.beta.]]- (9CI)
(CA INDEX NAME)

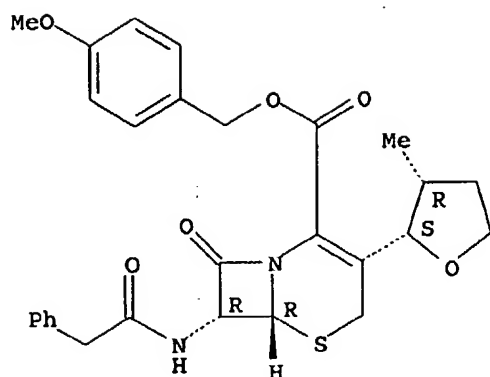
Absolute stereochemistry.



RN 141194-83-4 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-(tetrahydro-3-methyl-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-[3(2S*,3R*),6.alpha.,7.beta.]]- (9CI)
(CA INDEX NAME)

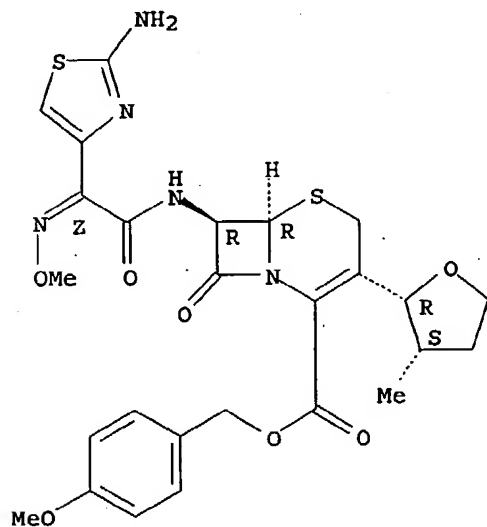
Absolute stereochemistry.



RN 141194-85-6 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl) (methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-3-methyl-2-furanyl)-, (4-methoxyphenyl)methyl ester, [6R-[3(2R*,3S*),6.alpha.,7.beta.(Z)]]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.



L6 ANSWER 7 OF 7 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1992:128438 HCAPLUS

DOCUMENT NUMBER: 116:128438

TITLE: Penicillin-cephalosporin conversion. XV. A new short-cut route to 3-norcephalosporins

AUTHOR(S): Tanaka, Hideo; Kameyama, Yutaka; Sumida, Shinichi; Yamada, Takae; Tokumaru, Yoshihisa; Shiro, Takashi; Sasaoka, Michio; Taniguchi, Mastoshi; Torii, Sigeru

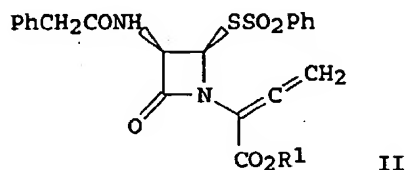
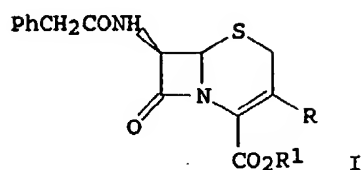
CORPORATE SOURCE: Fac. Eng., Okayama Univ., Okayama, 700, Japan

SOURCE: Synlett (1991), (12), 888-90

CODEN: SYNLES; ISSN: 0936-5214

Searched by Thom Larson, STIC, 308-7309

DOCUMENT TYPE: Journal
 LANGUAGE: English
 OTHER SOURCE(S): CASREACT 116:128438
 GI



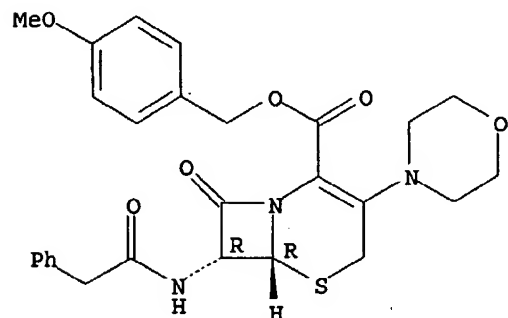
AB 3-Norcephems I (R = heterocyclic amino, SO₂Ph, Cl; R₁ = CH₂C₆H₄OMe-4, CHPh₂) were prepd. starting from penicillin G through a new shortcut involving Michael addn. of amines, azide or thiols to allenic esters II and sequential ring closure to the six-membered ring.

IT 139472-61-0P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of)

RN 139472-61-0 HCAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
 3-(4-morpholinyl)-8-oxo-7-[(phenylacetyl)amino]-, (4-methoxyphenyl)methyl
 ester, (6R-trans)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

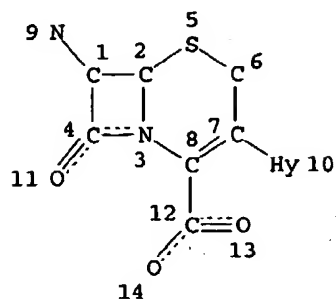


$$R_3 = \begin{pmatrix} -1 & -1 & 1 \end{pmatrix}^T_A$$

Page 1

L1

STR

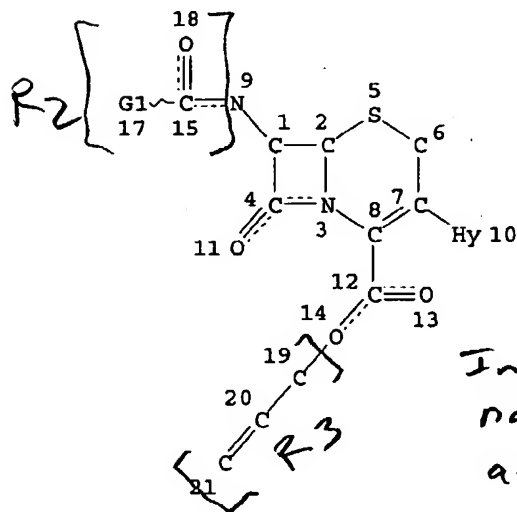


Same instnd search
as in previous
searches

ECOUNT IS E4 C E1 0 AT 10

NUMBER OF NODES IS 14

L5	STR
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Initially I defined R_3 more narrowly - but I didn't get any hits.

ECOUNT IS E4 C E1 O AT 10

NUMBER OF NODES IS 20

STEREO ATTRIBUTES: NONE

L7 4 SEA FILE=REGISTRY SUB=L2 SSS FUL L5

L8 4 SEA FILE=CAPLUS ABB=ON PLU=ON L7

*search L2 using structure
L5.**search CAPLUS with
answer set from
Registry*

=> D IBIB ABS HITSTR 1-4

L8 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:449689 CAPLUS

DOCUMENT NUMBER: 137:33162

TITLE: Process for the preparation of p-nitrobenzyl or allyl
esters of 3-cyclic-ether substituted cephalosporins
from trimethylphosphinic compounds via an
intramolecular Wittig reactionINVENTOR(S): Colberg, Juan Carlos; Tucker, John Lloyd; Zenoni,
Maurizio; Fogliato, Giovanni; Donadelli, Alessandro

PATENT ASSIGNEE(S): Pfizer Products Inc., USA

SOURCE: PCT Int. Appl., 47 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002046199	A1	20020613	WO 2001-IB2181	20011119
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			

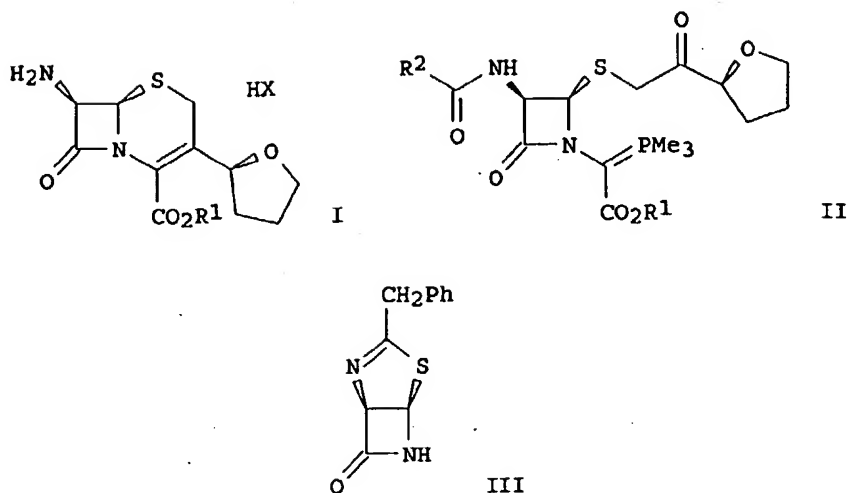
US 2002099205 A1 20020725 US 2001-6579 20011204

PRIORITY APPLN. INFO.: US 2000-251018P P 20001204

OTHER SOURCE(S): CASREACT 137:33162; MARPAT 137:33162

GI

*Applicants'
work.*



AB A process for the prepn. of I (R₁ = p-nitrobenzyl, allyl; X = halo) via an intramol. Wittig reaction of II (R₁ = p-nitrobenzyl, allyl; R₂ = C1-6-alkyl, C6-10-aryl, C6-10-aryl-C1-6-alkyl, dithianyl) to prep. 3-cyclic-ether substituted derivs. of cephalosporins is described. Thus, III was treated with p-nitrobenzyl glyoxylate monohydrate followed by redn. of the intermediate with NaBH₄. The resulting hydroxy compd. was treated with p-toluenesulfonic acid followed by addn. of (S)-1-(tetrahydro-2-furanyl)ethanone, addn. of thionyl chloride, and finally trimethylphosphine to give the desired intermediate II (R₁ = p-nitrobenzyl, R₂ = PhCH₂). Cyclization of II via an intramol. Wittig reaction was accomplished by refluxing for 16 h in THF. Addn. of phosphorus pentachloride and .alpha.-picoline in dichloromethane gave the free amine of I (R₁ = p-nitrobenzyl).

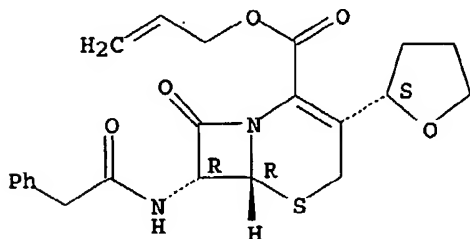
IT 436100-76-4P

RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(process for the prepn. of p-nitrobenzyl or allyl esters of 3-cyclic-ether substituted cephalosporins from trimethylphosphinic compds. via an intramol. Wittig reaction)

RN 436100-76-4 CAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid, 8-oxo-7-[(phenylacetyl)amino]-3-[(2S)-tetrahydro-2-furanyl]-, 2-propenyl ester, (6R,7R)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT:

6

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS

Searched by Thom Larson, STIC, 308-7309

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:449688 CAPLUS

DOCUMENT NUMBER: 137:33161

TITLE: Coupling process and intermediates useful for
preparing cephalosporinsINVENTOR(S): Colberg, Juan Carlos; Donadelli, Alessandro; Fogliato,
Giovanni; Zenoni, Maurizio

PATENT ASSIGNEE(S): Pfizer Products Inc., USA

SOURCE: PCT Int. Appl., 33 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002046198	A1	20020613	WO 2001-IB2225	20011122
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			

PRIORITY APPLN. INFO.: US 2000-251014P P 20001204

OTHER SOURCE(S): MARPAT 137:33161

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB This invention relates to a novel process for the prepn. of 3-cyclic-ether-substituted cephalosporins, such as I [CO₂R₁ = carboxylic acid or a carboxylate salt; A₁ = aryl, heteroaryl, heterocyclyl; A₂ = H, alkyl, cycloalkyl, aryl, etc.], via amidation reactions. Thus, cephalosporin II was prepd. in 80% yield by amidation of amine III with the acid anhydride of acid IV using O,O-di-Et hydrogenphosphorothioate in a Me₂CO/H₂O soln.

IT 436100-70-8P 436100-76-4P

RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(process for the prepn. of intermediates via amidation which are useful for prepg. cephalosporins)

RN 436100-70-8 CAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2Z)-(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-[(2S)-tetrahydro-2-furanyl]-, 2-propenyl ester, (6R,7R)-, monobenzenesulfinate (9CI) (CA INDEX NAME)

CM 1

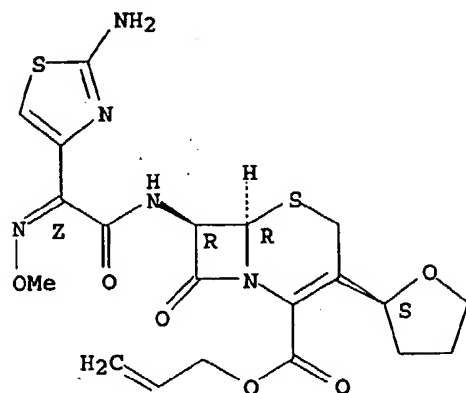
CRN 436100-69-5

CMF C20 H23 N5 O6 S2

Searched by Thom Larson, STIC, 308-7309

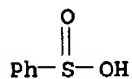
APP 1/2/02
work.

Absolute stereochemistry.
Double bond geometry as shown.



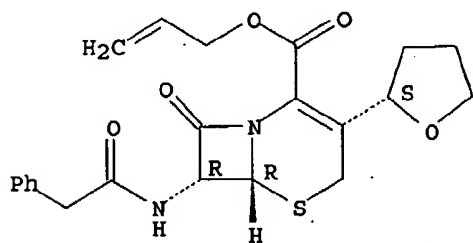
CM 2

CRN 618-41-7
CMF C6 H6 O2 S



RN 436100-76-4 CAPLUS
CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
8-oxo-7-[(phenylacetyl)amino]-3-[(2S)-tetrahydro-2-furanyl]-, 2-propenyl
ester, (6R,7R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

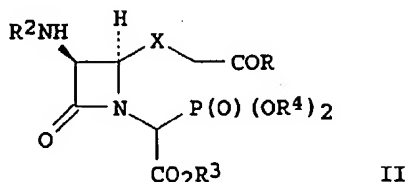
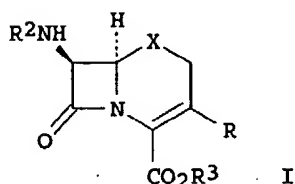
L8 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 1997:140924 CAPLUS
DOCUMENT NUMBER: 126:144046
TITLE: Beta-lactam preparation
INVENTOR(S): Harris, Michael Anthony; Saunders, Richard Neville
PATENT ASSIGNEE(S): Pfizer Limited, UK

Searched by Thom Larson, STIC, 308-7309

SOURCE: Brit. UK Pat. Appl., 15 pp.
 CODEN: BAXXDU
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
GB 2300856	A1	19961120	GB 1995-10126	19950516

OTHER SOURCE(S): CASREACT 126:144046; MARPAT 126:144046
 GI



AB Title compds. I [R = substituent; R1 = H, OMe, NHCHO; R2 = acyl; CO2R3 = CO2H, CO2-; R3 = protecting group; X = S, SO, SO2, O, CH2] are prepd. by base-induced cyclization of an azetidinone II [R4 = alkyl, aryl]. II are prepd. from the halide and P(OR4)3. Thus, 4-methoxybenzyl (2RS)-2-hydroxy-2-[(3R)(4R)-3-phenylacetamido-4-[(RS)-2-tetrahydrofuryl]carbonylmethylthio]azetidin-2-on-1-ylacetate was converted to the chloride and then to the phosphonate which was cyclized with NaH in PhMe to give 50% I [R = (RS)-2-tetrahydrofuryl, R1 = H, R2 = PhCH2CO, R3 = 4-MeC6H4CH2].

IT 141060-97-1P

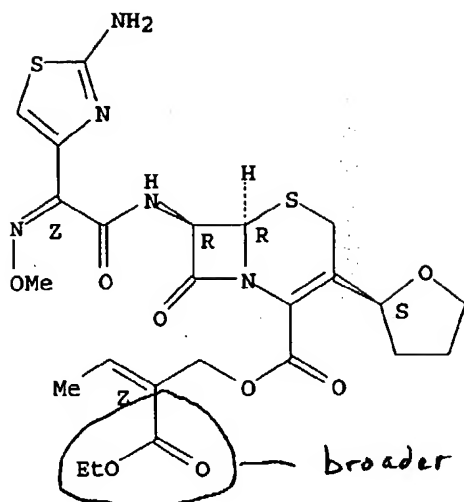
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(prepn. of cepheps by cyclization of azetidinyolphosphonoacetates with base)

RN 141060-97-1 CAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid, 7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-2-furanyl)-, 2-(ethoxycarbonyl)-2-butenyl ester, [6R-{2(Z),3(S*),6.alpha.,7.beta.(Z)}]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
 Double bond geometry as shown.



L8 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1992:255397 CAPLUS

DOCUMENT NUMBER: 116:255397

TITLE: Preparation of 3-tetrahydrofurylcephem-3-carboxylates and analogs as antibiotics

INVENTOR(S): Bateson, John Hargreaves; Burton, George; Fell, Stephen Christopher Martin

PATENT ASSIGNEE(S): Beecham Group PLC, UK

SOURCE: PCT Int. Appl., 147 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9201696	A1	19920206	WO 1991-GB1228	19910722
W: AU, CA, CS, FI, HU, JP, KR, NO, PL, US				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LU, NL, SE				
CA 2087967	AA	19920125	CA 1991-2087967	19910722
AU 9182224	A1	19920218	AU 1991-82224	19910722
AU 648329	B2	19940421		
ZA 9105725	A	19920624	ZA 1991-5725	19910722
EP 540609	A1	19930512	EP 1991-913583	19910722
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE				
HU 63628	A2	19930928	HU 1993-177	19910722
JP 05509305	T2	19931222	JP 1991-512368	19910722
JP 2851428	B2	19990127		
AT 185567	E	19991015	AT 1991-913583	19910722
ES 2137162	T3	19991216	ES 1991-913583	19910722
CN 1060469	A	19920422	CN 1991-105783	19910724
CN 1061046	B	20010124		
NO 9300226	A	19930323	NO 1993-226	19930122
US 6020329	A	20000201	US 1997-958864	19971020
CN 1223859	A	19990728	CN 1998-122407	19981114
US 6001997	A	19991214	US 1999-228138	19990111
US 6077952	A	20000620	US 1999-327667	19990608

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PRIORITY APPLN. INFO.:

GB 1990-16189	A 19900724
GB 1991-9540	A 19910502
WO 1991-GB1228	A 19910722
US 1993-934667	B1 19930122
US 1995-470786	B1 19950606
US 1997-958864	A1 19971020
US 1999-228138	A1 19990111

OTHER SOURCE(S): MARPAT 116:255397

GI For diagram(s), see printed CA Issue.

AB Title compds. (I; R1 = H, MeO, HCONH; R2 = acyl; R3 = H, neg. charge, carboxy-protective group; R4 = .ltoreq.4 substituents selected from alkyl, alkenyl, OH, halo, alkoxy, etc.; X = O, CH2, SOn; n= 0-2; m = 1, 2) were prepd. Thus, Na 2-(2-tritylaminothiazol-4-yl)-2-(Z)-trityloxyiminoacetate was condensed with tert-butyl (6R, 7R)-7-amino-3-[(R)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate to give, after deprotection, (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-hydroxyiminoacetamido]-3-[(RS)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylic acid which had MIC of 0.50 and 0.25 .mu.g/mL against Escherichia coli (NCTC 1048) and Staphylococcus aureus (Oxford), resp.

IT 141060-97-1P

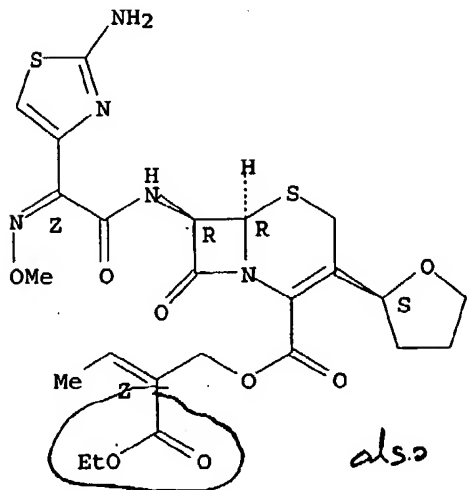
RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)
(prepn. of, as antibiotic)

RN 141060-97-1 CAPLUS

CN 5-Thia-1-azabicyclo[4.2.0]oct-2-ene-2-carboxylic acid,
7-[[[(2-amino-4-thiazolyl)(methoxyimino)acetyl]amino]-8-oxo-3-(tetrahydro-2-furanyl)-, 2-(ethoxycarbonyl)-2-butenyl ester, [6R-
[2(Z),3(S*),6.alpha.,7.beta.(Z)]]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.



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